

STIC Search Report Biotech-Chem Library

STIC Database Tracking Number: 12798

TO: Ben Sackey

Location: rem/5b31/5c18

Art Unit: 1626

Monday, July 26, 2004

Case Serial Number: 10/618044

From: Noble Jarrell

Location: Biotech-Chem Library

Rem 1B71

Phone: 272-2556

Noble.jarrell@uspto.gov

Search Notes	Strategy of the strategy of th	





SEARCH REQUEST FORM

Scientific and Technical Information Center

	Requester's Full Name: BEN Art Unit: 1626 Phone Mail Box and Bldg/Room Location	SA CKEY Number 30 2-0704 n:16m 5 8 2 1 Res	Examiner #: 73489 Date: 7/36/04 Serial Number: 10/6/8,044 ults Format Preferred (circle): PAPER DISK E-MAIL				
	If more than one search is submitted, please prioritize searches in order of need.						
	Please provide a detailed statement of the Include the elected species or structures, k	search topic, and describe teywords, synonyms, acrost that may have a special m	as specifically as possible the subject matter to be searched. nyms, and registry numbers, and combine with the concept or eaning. Give examples or relevant citations, authors, etc, if				
	Title of Invention: Wicyclic	Protein	Kigase Libitors				
	Inventors (please provide full names):	Dan M. Bay	Kinase Inhibitors				
	Earliest Priority Filing Date:						
	For Sequence Searches Only Please include	de all pertinent information	 (parent, child, divisional, or issued patent numbers) along with the				
	appropriate serial number. (CH2) CN	Elect 8-(4-cl lingl)e nitil	ed species is inlove-5-methoxy-2-methylonilino)-3-22-(4-morpho ehyl]-3H-imida-2024,5-g]quindine-7-(4-bo				
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	Searcher: Noble	NA Sequence (#)	STN 639				
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	Date Searcher Picked Up: 126104	Bibliographic	Dr.Link				
	Date Completed: 16009	Litigation	Lexis/Nexis				
	Searcher Prep & Review Time:	Fulltext	Sequence Systems				
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(FILE 'HOME' ENTERED AT 08:57:21 ON 26 JUL 2004)

FILE MCAPLUS: ENTERED AT 08:57:30 ON 26 JUL 2004
L1 1 UB2064110762/DN

FILE 'REGISTRY' ENTERED AT 08:57:45 ON 26 JUL 2004

FILE 'HCAPLUS' ENTERED AT 08:57:50 ON 26 JUL 2004 L2 TRA L1 1- RN : 291 TERMS

FILE 'REGISTRY' ENTERED AT 08:57:51 ON 26 JUL 2004
L3 291 SEA D2

FILE 'REGISTRY' ENTERED AT 08:57:54 ON 26 JUL 2004

FILE ENTERED AT 08:57:56 ON 26 JUL 2004 L4 US2004110762/PN

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FILE COVERS 1907 - 26 Jul 2004 VOL 141 ISS 5 FILE LAST UPDATED: 25 Jul 2004 (20040725/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

'OBI' IS DEFAULT SEARCH FIELD FOR 'HCAPLUS' FILE

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- L1 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2004 ACS on STN
- AN 2001:906207 HCAPLUS
- DN 136:37618
- ED Entered STN: 16 Dec 2001
- TI Preparation of substituted aromatic tricyclic compounds containing nicotinonitrile rings as protein kinase inhibitors
- IN * Berger, Dan M.; Dutia, Minu D.; Demorin, Frenel F.; Boschelli, Diane H.;
 Powell, Dennis W.; Tsou, Hwei-ru; Wissner, Allan; Zhang, Nan; Ye, Fei; Wu,
 Biqi
- PA American Home Products Corporation, USA; Wyeth
- SO U.S. Pat. Appl. Publ., 107 pp. CODEN: USXXCO
- DT Patent

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LA English
IC ICM A6
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C ICM A61K031-5377

ICS A61K031-496; A61K031-4738; C07D491-02

NCL 514232800

CC 28-13 (Heterocyclic Compounds (More Than One Hetero Atom))

Section cross-reference(s): 1

FAN.CNT 1

	0111 1				
	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	US 2001051620	A1	20011213	US 2000-751274	20001229
	US 6638929	B2	20031028		
	US 2004110762	A1	20040610	US 2003-618044	20030710 <
PRAI	US 1999-240905P	P	19991229		
	US 2000-751274	A3	20001229		
OS	MARPAT 136:37618				
GI					

The title compds. I [Ar = (un)substituted cycloalkyl, pyridyl, pyrimidinyl, etc.; n = 0-1; X = NH, O, S, NR; R = alkyl; Y, Z = both carbon or N; A = (un)substituted benzo, pyrido, pyrimido, etc.] which are useful as inhibitors of protein tyrosine kinase and are antiproliferative agents, were prepared E.g., a 3-step synthesis of II which showed IC50 of 0.005 .mu.M against EGF-R kinase (recombinant enzyme), was given.

arom tricyclic compd prepn protein kinase inhibitor; EGF receptor kinase inhibitor arom tricyclic compd prepn; antitumor arom tricyclic compd prepn; KDR kinase inhibitor arom tricyclic compd prepn; mitogen activated protein kinase inhibitor arom tricyclic compd prepn; src kinase inhibitor arom tricyclic compd prepn

IT Antitumor agents

(preparation of substituted aromatic tricyclic compds. containing nicotinonitrile

rings as protein kinase inhibitors)

TT 79079-06-4, EGF receptor kinase 139691-76-2, Raf kinase 141349-89-5, Src kinase 142243-02-5, Mitogen activated protein kinase 150977-45-0 RL: BSU (Biological study, unclassified); BIOL (Biological study)

(preparation of substituted aromatic tricyclic compds. containing nicotinonitrile

rings as protein kinase inhibitors)

IT	263149-40-2P	348617-29-8P	348617-39-0P	348617-40-3P	348617-42-5P
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	348617-64-1P	348617-89-0P	348617-94-7P	348617-95-8P	348618-04 - 2P
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	348618-65-5P	348618-81-5P	348619-28-3P		

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    preparation); THU (Therapeutic use); BIOL (Biological study); PREP
     (Preparation); RACT (Reactant or reagent); USES (Uses)
        (preparation of substituted aromatic tricyclic compds. containing
nicotinonitrile
       rings as protein kinase inhibitors)
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                                                  348617-26-5P
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     348617-17-4P
                    348617-19-6P
                                                                 348617-44-7P
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     RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU
     (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
     (Uses)
        (preparation of substituted aromatic tricyclic compds. containing
nicotinonitrile
       rings as protein kinase inhibitors)
     79-10-7, Acrylic acid, reactions 90-05-1, Guaiacol
                                                            94-05-3, Ethyl
TΤ
     (ethoxymethylene))cyanoacetate 105-34-0, Methyl cyanoacetate
                                                                      108-01-0,
     2-(Dimethylamino)ethanol 109-01-3, 1-Methylpiperazine
                                                               110-91-8,
                           139-59-3, 4-Phenoxyaniline
     Morpholine, reactions
                                                         288-36-8,
                        348-62-9, 4-Chloro-2-fluorophenol
                                                            367-21-5,
     1H-1,2,3-Triazole
     3-Chloro-4-fluoroaniline
                              504-88-1, 3-Nitropropionic acid
                                                                 540-88-5,
     tert-Butyl acetate 554-00-7, 2,4-Dichloroaniline
                                                         591-19-5,
     3-Bromoaniline 622-40-2, 4-(2-Hydroxyethyl) morpholine
                                                               632-02-0,
     3-Chloropropyl p-toluenesulfonate 814-68-6, Acryloyl chloride
     873-38-1, 2-Bromo-4-chloroaniline
                                         882-33-7, Phenyl disulfide
     1142-19-4, 4,4'-Dichlorodiphenyl disulfide 2038-03-1,
     4-(2-Aminoethyl)morpholine 2835-95-2, 5-Amino-o-cresol
                                                               4637-24-5
     5335-29-5, 3-Chloro-4-phenoxyaniline 5959-52-4, 3-Amino-2-naphthoic acid
     20357-25-9, 6-Nitroveratraldehyde
                                       24313-88-0, 3,4,5-Trimethoxyaniline
     33693-48-0, 4-Benzyloxy-3-methoxybenzyl alcohol 34674-75-4
                                                                   35212-85-2,
     Methyl 3-aminobenzo[b]thiophene-2-carboxylate 39786-35-1, Ethyl
     3-amino-2-benzo[b] furancarboxylate 43073-44-5, 6,7-Dimethoxy-2,3-
     naphthalenedicarboxylic anhydride 50868-72-9, 5-Methoxy-2-methylaniline
     57946-56-2, 4-Chloro-2-fluoroaniline 59404-86-3, 4-Benzyloxy-3-
                    59922-33-7
                                62492-42-6
                                               63224-35-1
                                                            76513-69-4,
     chloroaniline
     2-(Trimethylsilyl)ethoxymethyl chloride
                                               76878-17-6
                                                            85006-21-9,
                                               98404-04-7, 2-Chloro-4-fluoro-5-
     2-Chloro-5-methoxyaniline hydrochloride
                      98446-49-2, 2,4-Dichloro-5-methoxyaniline
                                                                 131775-97-8
     methoxyaniline
                  133303-88-5
                                 204915-71-9, 4-(2-Chloroethoxy)-3-
     133088-44-5
     methoxybenzaldehyde 348619-47-6
     RL: RCT (Reactant); RACT (Reactant or reagent)
```

```
(preparation of substituted aromatic tricyclic compds. containing
nicotinonitrile
       rings as protein kinase inhibitors)
                                         53544-07-3P 53815-60-4P
    3590-37-2P, Ethyl 3-nitropropionate
                                  309269-57-6P
                                                348617-15-2P
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    RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
     (Reactant or reagent)
        (preparation of substituted aromatic tricyclic compds. containing
nicotinonitrile
       rings as protein kinase inhibitors)
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     treatment, inhibition and eradication of neoplasms, polycystic kidney
     disease and colonic polyps.
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     H; WISSNER, A; WU, B; YE, F; ZHANG, N
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     US 2000-751274 20001229; CN 1437584 A CN 2000-819209 20001229; US
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     US 2003-618044
                          20030710
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AΒ

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WO 200147892 A UPAB: 20011005
NOVELTY - Tricyclic heterocycles (I) are new.
              DETAILED DESCRIPTION - Tricyclic heterocycles of formula (I) and
their salts are new.
              ring system = fused 5,6,6-tricycle, 6,6,6-tricycle, 6,5,6-tricycle,
or 6,6,6-tricycle;
              Ar = 3-7C cycloalkyl (optionally substituted by 1 or more 1-6C
alkyl), or pyridinyl, pyrimidinyl or phenyl (all optionally substituted),
8-12 membered bicyclic aryl or bicyclic heteroaryl containing 1-4 N, O or
S (both optionally substituted), or -A'-T-L;
              A' = pyridinyl, pyrimidinyl or phenyl (all optionally substituted);
T = -NH(CH2)m-, -O(CH2)m-, -S(CH2)m-, -NR(CH2)m-, -(CH2)m-, -(CH
-(CH2) mCO-, -(CH2) mSO-, -(CH2) mSO2- or -(CH2) mNR-;
              L = phenyl (optionally substituted), or 5-6 membered heteroaryl ring
containing 1-3 N, O or S and optionally substituted;
m = 0-3;
n = 0-1;
              X = NH, O, S or NR;
R = 1-6C \text{ alkyl};
              Y, Z = C or N; or
              Y = N, O or S; and
Z = bond; or
Y = bond; and
Z = N, O or S;
              ring A = a group of formula (a) - (1);
B = C \text{ or } N;
              D, E = C, N, O or S;
              dotted line = optional double bond;
              R1-R4 = absent, H, OH, halo, amino, hydroxyamino, CF3, CF3O, SH, 1-6C
alkyl, 3-8C cycloalkyl, 2-6C alkenyl, 2-6C alkynyl, 2-6C alkenyloxy, 2-6C
alkynyloxy, 1-6C hydroxyalkyl, 1-6C mercaptoalkyl, halomethyl, 2-7C
alkoxymethyl, 1-6C alkoxy, 3-8C cycloalkoxy, 1-6C alkylthio, 3-8C
cycloalkylthio, 1-6C alkylsulfinyl, 1-6C alkylsulfonyl, 1-6C
alkylsulfonamido, 2-6C alkenylsulfonamido, 2-6C alkynylsulfonamido, CN,
NO2, carboxy, 2-7C alkoxycarbonyl, 2-7C alkanoyl, 3-7C alkenoyl, 4-12C
N-alkyl-N-alkenylamino, 6-12C dialkenylamino, phenylamino, benzylamino,
phenoxy, phenyl, thiophenoxy, benzyl, 1-6C alkylamino, 2-7C alkanoyloxy,
3-8C alkenoyloxy, 3-8C alkynoyloxy, carbamoyl, 2-7C N-alkylcarbamoyl,
3-13C N,N-dialkylcarbamoyl, 2-12C dialkylamino, 2-7C alkanoyloxymethyl,
2-7C alkenoyloxymethyl, 2-7C alkynoyloxymethyl, azido, benzoyl, 2-7C
carboxyalkyl, 3-8C carboxyalkoxyalkyl, R8R9-CH-M-(C(R6)2)k-X-,
R7 - (C(R6)2)g-X-, R7 - (C(R6)2)p-M-(C(R6)2)k-X-, Het-(C(R6)2)q-W-(C(R6)2)k-X-
  , Ph-(C(R6)2)q-W-(C(R6)2)k-X-, R5-CONH(CH2)q-, R5-C equivalent to
 {\tt C-CONH\,(CH2)\,q^-\,,\quad (R5)\,2C=C\,(CN)\,\,(CH2)\,q^-\,,\quad (R5)\,2C=C\,(R5)\,CONH\,(CH2)\,q^-\,,} 
  (\text{R5}) \ 2\text{C} = \text{C} \ (\text{R5}) \ \text{SO2NH} \ (\text{CH2}) \ q^- \ , \quad \text{R5OC} \ (\text{O}) \ \text{NH} \ (\text{CH2}) \ q^- \ , \quad \text{R5NHC} \ (\text{O}) \ \text{NH} \ (\text{CH2}) \ q^- \ , 
  (\text{R5}) \, 2\text{NC} \, (\text{O}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{O}) \, \text{O} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, , \quad \text{R5HNC} \, (\text{S}) \, \text{NH} \, (\text{CH2}) \, q\text{-} \, 
 (R5) 2NC(S) NH(CH2) q-, (R5) 2NC(O)O(CH2) q-, or a group of formula (m) or (n);
               R5 = H, 1-6C alkyl, 1-6C aminoalkyl, 2-9C N-alkylaminoalkyl, 3-12C
N, N-dialkylaminoalkyl, 4-12C N-cycloalkylaminoalkyl, 5-18C
N-cycloalkyl-N-alkylaminoalkyl, 7-18C N, N-dicycloalkylaminoalkyl,
 morpholino-N-(1-6C alkyl), piperidino-N-(1-6C alkyl), N-(1-6C
 alkyl)-piperazino-N-(1-6C alkyl), 3-11C azacycloalkyl-N-alkyl, 1-6C
hydroxyalkyl, 2-8C alkoxyalkyl or phenyl;
              X = (CH2)m, O, S or NR6;
               R7 = N(R6)2, OR6, J, N(R6)3 + or NR6(OR6);
               M = NR6, O, S, N-((C(R6)2)pN(R6)2) or N-((C(R6)2)p-OR6);
               W = NR6, O, S or bond;
               Het = morpholine, thiomorpholine, thiomorpholine-S-oxide,
 thiomorpholine-S,S-dioxide, piperidine, pyrrolidine, aziridine, pyridine,
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imidazole, 1,2,3-triazole, 1,2,4-triazole, thiazole, thiazolidine,
tetrazole, piperazine, furan, thiophene, tetrahydrothiophene,
tetrahydrofuran, dioxane, 1,3-dioxolane pyrrole or tetrahydropyran (all
optionally C-substituted, and optionally N-substituted by 1-2 NR6);
     Ph = optionally substituted phenyl;
     R6 = H, 1-6C alkyl, 2-6C alkenyl, 2-6C alkynyl, 1-6C cycloalkyl, 2-7C
alkanoyl, 2-7C carbamoylalkyl, 1-6C hydroxyalkyl, 3-6C hydroxycycloalkyl
or 2-7C carboxyalkyl, or optionally substituted phenyl;
     R8, R9 = N-((C(R6)2)pN(R6)2) \text{ or } N-((C(R6)2)p-OR6);
     J = H, Cl, F or Br;
q = 1-6;
k = 0-4;
p = 2-4;
q = 0-4;
r = 1-4; and
s = 1-6;
provided that:
(1) when ring A = group (1), at least 1 bond between E and B or B and D is a double bond; at least 1 of E, B or D is not C; only 1 of E, B or D
= 0 or S; and the adjacent atoms to 0 or S are C;
     (2) when R5 is bound to an N atom, the resulting structures are not
N-C-N or O-C-N radicals; and when R5 is bound to an O atom, the resulting
structure is not a -N-C-O- radical;
     (3) when R6 = 2-7C alkenyl or 2-7C alkynyl, the alkenyl and alkynyl
moieties are bound to N or O through a saturated C atom in the alkenyl or
alkynyl chain;
     (4) when X = NR6 and R7 = N(R6)2, N(R6)3+ or NR6(OR6), then g = 2-6;
     (5) when M = 0 or S and R7 = 0R6, then p = 1-4;
     (6) when X = NR6, O or S, then k = 2-4;
     (7) when X = 0 or S and M or W = 0 or S, then k = 1-4;
     (8) when W is not a bond with Het bonded through an N atom, then q =
2-4; and
     (9) when W = bond with Het bonded through an N atom and X = NR6, O or
S, then k = 2-4.
     An INDEPENDENT CLAIM is also included for the preparation of (I) via
14 different reaction pathways.
     ACTIVITY - Cytostatic; Nephrotropic; Gastrointestinal; Hepatotropic;
Dermatological.
     MECHANISM OF ACTION - MAPK kinase inhibitor; RAF kinase inhibitor;
SRC kinase inhibitor; ECK/LERK-1 kinase inhibitor; VEGF/KDR kinase
inhibitor
     In an assay to measure inhibition of cancer cell growth (see Skehan
et. al., J. Natl. Canc. Inst., 82,1107-1112 (1990)), 4-(2,4-
dichloroanilino) -8-nitro(1) benzothieno(3,2-b) pyridine-3-carbonitrile (Ib)
displayed IC50 values of 0.46, 0.41, 0.59 and 0.67 micro g/ml against
MDA-MB-435, A431, SK-BR3 and SW620 cell lines respectively.
     USE - Compounds (I) are useful for treating, eradicating or
inhibiting the growth of neoplasms, especially breast, kidney, bladder,
mouth, larynx, esophagus, stomach, colon, ovary, lung, pancreas, liver,
prostate and skin, and especially neoplasms that express EGFR or erbB2
(Her2). (I) are also useful in treating, eradicating or inhibiting
polycystic kidney disease and colonic polyps, and for inhibiting the
biological effects of a deregulated protein kinase (all claimed).
Dwg.0/0
CPI
AB; GI; DCN
CPI: B05-B01B; B06-H; B10-B01A; B10-B02A; B10-G02; B14-D06; B14-H01B;
     B14-N10
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FS

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MC

=> b home

FILE 'HOME' ENTERED AT 08:58:36 ON 26 JUL 2004

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=> b reg

FILE 'REGISTRY ENTERED AT 10:03:41 ON 26 JUL 2004
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
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Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 23 JUL 2004 HIGHEST RN 715654-51-6 DICTIONARY FILE UPDATES: 23 JUL 2004 HIGHEST RN 715654-51-6

TSCA INFORMATION NOW CURRENT THROUGH MAY 21, 2004

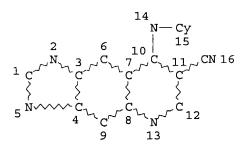
Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at: http://www.cas.org/ONLINE/DBSS/registryss.html

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L7



STR

NODE ATTRIBUTES:
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES: RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 16

STEREO ATTRIBUTES: NONE

19 20 SEA FILE=REGISTRY SECTION 12

100.0% PROCESSED 501 ITERATIONS SEARCH TIME: 00.00.01

20 ANSWERS

- a lot le tot

- L9 ANSWER 1 OF 20 REGISTRY COPYRIGHT 2004 ACS on STN
- RN 500023-80-3 REGISTRY
- CN 3H-Imidazo[4,5-g]quinoline-7-carbonitrile, 3-[2-(4-morpholinyl)ethyl]-8-[(3,4,5-trimethoxyphenyl)amino]- (9CI) (CA INDEX NAME)

FS 3D CONCORD

MF C26 H28 N6 O4

SR CA

LC STN Files: CA, CAPLUS, CASREACT DT.CA CAplus document type: Journal

RL.NP Roles from non-patents: BIOL (Biological study); PREP (Preparation)

$$\begin{array}{c} \text{OMe} \\ \text{MeO} \\ \text{OMe} \\ \text{NH} \\ \text{CN} \\ \\ \text{O} \\ \text{N} \end{array}$$

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1907 TO DATE)

1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L9 ANSWER 2 OF 20 REGISTRY COPYRIGHT 2004 ACS on STN

RN 500023-79-0 REGISTRY

CN 3H-Imidazo[4,5-g]quinoline-7-carbonitrile, 8-[(2-bromo-4-chloro-5-methoxyphenyl)amino]-3-[2-(4-morpholinyl)ethyl]- (9CI) (CA INDEX NAME)

FS 3D CONCORD

MF C24 H22 Br Cl N6 O2

SR CA

LC STN Files: CA, CAPLUS, CASREACT

DT.CA CAplus document type: Journal

RL.NP Roles from non-patents: BIOL (Biological study); PREP (Preparation)

$$\begin{array}{c|c} & \text{Cl} \\ & \text{MeO} \\ & \text{NH} \\ & \text{CN} \\ & \text{N} \\ & \text{CN} \\ & \text{N} \\ & \text{CN} \\ & \text{N} \\ & \text{N$$

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1907 TO DATE)
1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L9 ANSWER 3 OF 20 REGISTRY COPYRIGHT 2004 ACS on STN

RN 500023-78-9 REGISTRY

CN 3H-Imidazo[4,5-g]quinoline-7-carbonitrile, 8-[(5-methoxy-2-methylphenyl)amino]-3-[2-(4-morpholinyl)ethyl]- (9CI) (CA INDEX NAME)

FS 3D CONCORD

MF C25 H26 N6 O2

SR CA

LC STN Files: CA, CAPLUS, CASREACT DT.CA CAplus document type: Journal

RL.NP Roles from non-patents: BIOL (Biological study); PREP (Preparation)

$$\begin{array}{c|c} & \text{MeO} \\ & \text{Me} \\ & \text{NH} \\ & \text{CN} \\ & \text{O} \\ & \text{N} \\ & \text{CH}_2 - \text{CH}_2 \\ & \text{N} \\ & \text$$

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1907 TO DATE)

1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L9 ANSWER 4 OF 20 REGISTRY COPYRIGHT 2004 ACS on STN

RN 500023-77-8 REGISTRY

CN 3H-Imidazo[4,5-g]quinoline-7-carbonitrile, 8-[(2-chloro-5-methoxyphenyl)amino]-3-[2-(4-morpholinyl)ethyl]- (9CI) (CA INDEX NAME)

FS 3D CONCORD

MF C24 H23 Cl N6 O2

SR CA

LC STN Files: CA, CAPLUS, CASREACT

DT.CA CAplus document type: Journal

RL.NP Roles from non-patents: BIOL (Biological study); PREP (Preparation)

$$\begin{array}{c|c} & \text{MeO} \\ & \text{NH} \\ & \text{CN} \\ \\ & \text{O} \end{array}$$

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- 1 REFERENCES IN FILE CA (1907 TO DATE)
- 1 REFERENCES IN FILE CAPLUS (1907 TO DATE)
- L9 ANSWER 5 OF 20 REGISTRY COPYRIGHT 2004 ACS on STN
- RN 348619-42-1 REGISTRY
- CN Formamide, N-(7-cyano-1H-imidazo[4,5-g]quinolin-8-yl)-N-(4-phenoxyphenyl)(9CI) (CA INDEX NAME)
- FS 3D CONCORD
- MF C24 H15 N5 O2
- SR CA
- LC STN Files: CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL
- DT.CA CAplus document type: Patent
- RL.P Roles from patents: PREP (Preparation); RACT (Reactant or reagent)

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- 2 REFERENCES IN FILE CA (1907 TO DATE)
- 2 REFERENCES IN FILE CAPLUS (1907 TO DATE)
- L9 ANSWER 6 OF 20 REGISTRY COPYRIGHT 2004 ACS on STN
- RN 348619-39-6 REGISTRY
- CN Formamide, N-(7-cyano-1H-imidazo[4,5-g]quinolin-8-yl)-N-(3,4,5-trimethoxyphenyl)- (9CI) (CA INDEX NAME)
- FS 3D CONCORD
- MF C21 H17 N5 O4

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPATZ, USPATFULL

DT.CA CAplus document type: Patent

RL.P Roles from patents: PREP (Preparation); RACT (Reactant or reagent)

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

2 REFERENCES IN FILE CA (1907 TO DATE)

2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L9 ANSWER 7 OF 20 REGISTRY COPYRIGHT 2004 ACS on STN

RN 348617-72-1 REGISTRY

CN 3H-Imidazo[4,5-g]quinoline-7-carbonitrile, 3-[2-(4-morpholinyl)ethyl]-8-[(4-phenoxyphenyl)amino]- (9CI) (CA INDEX NAME)

FS 3D CONCORD

MF C29 H26 N6 O2

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPAT7LL

DT.CA CAplus document type: Patent

RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

$$\begin{array}{c|c} & & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & & & \\ & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & & \\ & &$$

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

2 REFERENCES IN FILE CA (1907 TO DATE)

Searched by Noble Jarrell

2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L9 ANSWER 8 OF 20 REGISTRY COPYRIGHT 2004 ACS on STN

RN 348617-71-0 REGISTRY

CN 3H-Imidazo[4,5-g]quinoline-7-carbonitrile, 8-[(4-chloro-5-methoxy-2-methylphenyl)amino]-3-[2-(4-morpholinyl)ethyl]- (9CI) (CA INDEX NAME)

FS 3D CONCORD

MF C25 H25 Cl N6 O2

SR CA

LC STN Files: CA, CAPLUS, CASREACT, TOXCENTER, USPAT7LLL

DT.CA CAplus document type: Journal; Patent

RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

RL.NP Roles from non-patents: BIOL (Biological study); PREP (Preparation)

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- 3 REFERENCES IN FILE CA (1907 TO DATE)
- 3 REFERENCES IN FILE CAPLUS (1907 TO DATE)
- L9 ANSWER 9 OF 20 REGISTRY COPYRIGHT 2004 ACS on STN

RN 348617-62-9 REGISTRY

CN lH-Imidazo[4,5-g]quinoline-7-carbonitrile, 2-(4-morpholinylmethyl)-8-[(3,4,5-trimethoxyphenyl)amino]- (9CI) (CA INDEX NAME)

FS 3D CONCORD

DR 500023-76-7

MF C25 H26 N6 O4

SR CA

LC STN Files: CA, CAPLUS, CASREACT, TOXCENTER, USPAT2, USPATFULL

DT.CA CAplus document type: Journal; Patent

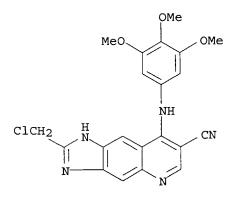
RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

RL.NP Roles from non-patents: BIOL (Biological study); PREP (Preparation)

$$\begin{array}{c|c} \text{OMe} \\ \text{MeO} \\ \text{OMe} \\ \text{OMe} \\ \text{NH} \\ \text{CN} \\ \text{OMe} \\ \text{NH} \\ \text{CN} \\ \text{OMe} \\ \text{OMe}$$

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- 3 REFERENCES IN FILE CA (1907 TO DATE)
- 3 REFERENCES IN FILE CAPLUS (1907 TO DATE)
- L9 ANSWER 10 OF 20 REGISTRY COPYRIGHT 2004 ACS on STN
- RN 348617-61-8 REGISTRY
- CN 1H-Imidazo[4,5-g]quinoline-7-carbonitrile, 2-(chloromethyl)-8-[(3,4,5-trimethoxyphenyl)amino]- (9CI) (CA INDEX NAME)
- FS 3D CONCORD
- MF C21 H18 Cl N5 O3
- SR CA
- LC STN Files: CA, CAPLUS, CASREACT, TOXCENTER, USPAT2, USPATFULL
- DT.CA CAplus document type: Journal; Patent
- RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)
- RL.NP Roles from non-patents: PREP (Preparation); RACT (Reactant or reagent)



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

- 3 REFERENCES IN FILE CA (1907 TO DATE)
- 3 REFERENCES IN FILE CAPLUS (1907 TO DATE)
- L9 ANSWER 11 OF 20 REGISTRY COPYRIGHT 2004 ACS on STN
- RN 348617-58-3 REGISTRY
- CN 1H-Imidazo[4,5-g]quinoline-7-carbonitrile, 8-[(4-phenoxyphenyl)amino]-

(9CI) (CA INDEX NAME)

FS 3D CONCORD

MF C23 H15 N5 O

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPAT7ULL

DT.CA CAplus document type: Patent
RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); USES
(Uses)

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

2 REFERENCES IN FILE CA (1907 TO DATE)
2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L9 ANSWER 12 OF 20 REGISTRY COPYRIGHT 2004 ACS on STN

RN 348617-56-1 REGISTRY

CN 1H-Imidazo[4,5-g]quinoline-7-carbonitrile, 8-[(3,4,5-trimethoxyphenyl)amino]- (9CI) (CA INDEX NAME)

FS 3D CONCORD

MF C20 H17 N5 O3

SR CA

LC STN Files: CA, CAPLUS, CASREACT, TOXCENTER, USPAT2, USPATFULL

DT.CA CAplus document type: Journal; Patent

RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

RL.NP Roles from non-patents: BIOL (Biological study); PREP (Preparation)

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

3 REFERENCES IN FILE CA (1907 TO DATE)
3 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L9 ANSWER 13 OF 20 REGISTRY COPYRIGHT 2004 ACS on STN

RN 348617-55-0 REGISTRY

CN 1H-Imidazo[4,5-g]quinoline-7-carbonitrile, 8-[(3-hydroxy-4-methylphenyl)amino]- (9CI) (CA INDEX NAME)

FS 3D CONCORD

MF C18 H13 N5 O

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPAT7, USPATFULL

DT.CA CAplus document type: Patent

RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

2 REFERENCES IN FILE CA (1907 TO DATE)

2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L9 ANSWER 14 OF 20 REGISTRY COPYRIGHT 2004 ACS on STN

RN 348617-54-9 REGISTRY

CN 1H-Imidazo[4,5-g]quinoline-7-carbonitrile, 8-[(2-chloro-5-methoxyphenyl)amino]- (9CI) (CA INDEX NAME)

FS 3D CONCORD

MF C18 H12 Cl N5 O

SR CA

LC STN Files: CA, CAPLUS, CASREACT, TOXCENTER, USPAT2, USPATFULL

DT.CA CAplus document type: Journal; Patent

RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

RL.NP Roles from non-patents: BIOL (Biological study); PREP (Preparation)

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

3 REFERENCES IN FILE CA (1907 TO DATE)

3 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L9 ANSWER 15 OF 20 REGISTRY COPYRIGHT 2004 ACS on STN

RN 348617-52-7 REGISTRY

CN 1H-Imidazo[4,5-g]quinoline-7-carbonitrile, 8-[(2-bromo-4-chloro-5-methoxyphenyl)amino]- (9CI) (CA INDEX NAME)

FS 3D CONCORD

MF C18 H11 Br Cl N5 O

SR CA

LC STN Files: CA, CAPLUS, CASREACT, TOXCENTER, USPATZ, USPATFULL

DT.CA CAplus document type: Journal; Patent

RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

RL.NP Roles from non-patents: BIOL (Biological study); PREP (Preparation)

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

3 REFERENCES IN FILE CA (1907 TO DATE)

3 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L9 ANSWER 16 OF 20 REGISTRY COPYRIGHT 2004 ACS on STN

RN 348617-51-6 REGISTRY

CN 1H-Imidazo[4,5-g]quinoline-7-carbonitrile, 8-[(2-bromo-4-chlorophenyl)amino]- (9CI) (CA INDEX NAME)

FS 3D CONCORD

MF C17 H9 Br Cl N5

SR CA

LC STN Files: CA, CAPLUS, CASREACT, TOXCENTER, USPAT7, USPATFULL

DT.CA CAplus document type: Journal; Patent

RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

RL.NP Roles from non-patents: BIOL (Biological study); PREP (Preparation)

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

3 REFERENCES IN FILE CA (1907 TO DATE)

3 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L9 ANSWER 17 OF 20 REGISTRY COPYRIGHT 2004 ACS on STN

RN 348617-50-5 REGISTRY

CN 1H-Imidazo[4,5-g]quinoline-7-carbonitrile, 8-[(3-bromophenyl)amino]- (9CI) (CA INDEX NAME)

FS 3D CONCORD

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SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPATZ, USPATFULL

DT.CA CAplus document type: Patent

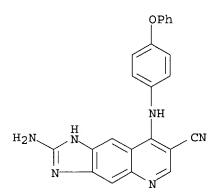
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PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

2 REFERENCES IN FILE CA (1907 TO DATE)

2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

- L9 ANSWER 18 OF 20 REGISTRY COPYRIGHT 2004 ACS on STN
- RN 348617-47-0 REGISTRY
- CN 1H-Imidazo[4,5-g]quinoline-7-carbonitrile, 2-amino-8-[(4-phenoxyphenyl)amino]-, monohydrochloride (9CI) (CA INDEX NAME)
- MF C23 H16 N6 O . Cl H
- SR CA
- LC STN Files: CA, CAPLUS, TOXCENTER, USPAT7LLL
- DT.CA CAplus document type: Patent
- RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)



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- 2 REFERENCES IN FILE CA (1907 TO DATE)
- 2 REFERENCES IN FILE CAPLUS (1907 TO DATE)
- L9 ANSWER 19 OF 20 REGISTRY COPYRIGHT 2004 ACS on STN
- RN 348617-46-9 REGISTRY
- CN 1H-Imidazo[4,5-g]quinoline-7-carbonitrile, 2-[[2-(4-morpholiny1)ethy1]amino]-8-[(3,4,5-trimethoxypheny1)amino]- (9CI) (CAINDEX NAME)
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- LC STN Files: CA, CAPLUS, CASREACT, TOXCENTER, USPATZ, USPATFULL
- DT.CA CAplus document type: Journal; Patent
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PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

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- RN 348617-44-7 REGISTRY
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- DT.CA CAplus document type: Journal; Patent
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Searched by Noble Jarrell

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FILE COVERS 1907 - 26 Jul 2004 VOL 141 ISS 5 FILE LAST UPDATED: 25 Jul 2004 (20040725/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

'OBT' IS DEFAULT SEARCH FIELD FOR 'HCAPLUS' FILE

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L26 ANSWER 1 OF 2 HCAPLUS COPYRIGHT 2004 ACS on STN

2001:906207 HCAPLUS ΔN

LI.50620 DN 136:37618 TI Pres Preparation of substituted aromatic tricyclic compounds containing nicotinonitrile rings as protein kinase inhibitors

Berger, Dan M.; Dutia, Minu D.; Demorin, Frenel F.; Boschelli, Diane H.; TN Powell, Dennis W.; Tsou, Hwei-ru; Wissner, Allan; Zhang, Nan; Ye, Fei; Wu, Biqi

American Home Products Corporation, USA; Wyeth PA

U.S. Pat. Appl. Publ., 107 pp. SO

CODEN: USXXCO

DT Patent

LΑ English

FAN.	FAN.CNT 1					
	PATENT NO.	KIND	DATE		APPLICATION NO.	DATE
ΡI	US 2001051620	A1	20011213		US 2000-751274	20001229 <
	US 6638929	B2	20031028			
	US 2004110762	A1	20040610		US 2003-618044	20030710 <
PRAI	US 1999-240905P	P	19991229	<		
	US 2000-751274	А3	20001229			
os	MARPAT 136:37618					
GI						

AB The title compds. I [Ar = (un)substituted cycloalkyl, pyridyl, pyrimidinyl, etc.; n = 0-1; X = NH, O, S, NR; R = alkyl; Y, Z = both carbon or N; A = (un)substituted benzo, pyrido, pyrimido, etc.] which are useful as inhibitors of protein tyrosine kinase and are antiproliferative agents, were prepared E.g., a 3-step synthesis of II which showed IC50 of 0.005 .mu.M against EGF-R kinase (recombinant enzyme), was given.

IT 348617-61-8P

RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(preparation of substituted aromatic tricyclic compds. containing nicotinonitrile

rings as protein kinase inhibitors)

IT 348617-44-7P 348617-46-9P 348617-47-0P

348617-50-5P 348617-51-6P 348617-52-7P

348617-54-9P 348617-55-0P 348617-56-1P

348617-58-3P 348617-62-9P 348617-71-0P

348617-72-1P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of substituted aromatic tricyclic compds. containing nicotinonitrile

rings as protein kinase inhibitors)

IT 348619-39-6P 348619-42-1P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of substituted aromatic tricyclic compds. containing nicotinonitrile

rings as protein kinase inhibitors)

L26 ANSWER 2 OF 2 HCAPLUS COPYRIGHT 2004 ACS on STN

AN 2001:489374 HCAPLUS

DN 135:92639

*TI Preparation of substituted aromatic tricyclic compounds containing nicotinonitrile rings as protein kinase inhibitors

MEN Berger, Dan M.; Dutia, Minu D.; Demorin, Frenel F.; Boschelli, Diane H.; Powell, Dennis W.; Tsou, Hwei-ru; Wissner, Allan; Zhang, Nan; Ye, Fei; Wu, Biqi

PA American Home Products Corp., USA

SO PCT Int. Appl., 377 pp.

CODEN: PIXXD2

DT Patent

LA English

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                     KIND DATE
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    WO 2000-US35616
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                           20001229
os
    MARPAT 135:92639
GΙ
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AB The title compds. I [Ar = (un)substituted cycloalkyl, pyridyl, pyrimidinyl, etc.; n = 0-1; X = NH, O, S, NR; R = alkyl; Y, Z = both carbon or N; A = (un)substituted benzo, pyrido, pyrimido, etc.] which are useful as inhibitors of protein tyrosine kinase and are antiproliferative agents, were prepared E.g., a 3-step synthesis of II which showed IC50 of 0.005 .mu.M against EGF-R kinase (recombinant enzyme), was given.

IT 348617-61-8P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(preparation of substituted aromatic tricyclic compds. containing nicotinonitrile

rings as protein kinase inhibitors)

IT 348617-44-7P 348617-46-9P 348617-47-0P 348617-50-5P 348617-51-6P 348617-52-7P 348617-54-9P 348617-55-0P 348617-56-1P

348617-58-3P 348617-62-9P 348617-71-0P

348617-72-1P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of substituted aromatic tricyclic compds. containing nicotinonitrile

rings as protein kinase inhibitors) TT 348619-39-6P 348619-42-1P RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent) (preparation of substituted aromatic tricyclic compds. containing nicotinonitrile rings as protein kinase inhibitors) THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT => b uspatall FILE 'USPATFUEL' ENTERED AT 10:05:40 ON 26 JUL 2004 CA INDEXING COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS) FILE ENTERED AT 10:05:40 ON 26 JUL 2004 CA INDEXING COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS) => d bib abs hitm 127 tot L27 ANSWER 1 OF 3 USPATFULL on STN AN 2004:145089 USPATFULL TITricyclic protein kinase inhibitors Berger, Dan M., New City, NY, UNITED STATES IN Dutia, Minu D., West Nyack, NY, UNITED STATES DeMorin, Frenel F., Nanuet, NY, UNITED STATES Boschelli, Diane H., New City, NY, UNITED STATES Powell, Dennis W., Westchester, NY, UNITED STATES Tsou, Hwei-Ru, New City, NY, UNITED STATES Wissner, Allan, Ardsley, NY, UNITED STATES Zhang, Nan, Eastchester, NY, UNITED STATES Ye, Fei, Nanuet, NY, UNITED STATES Wu, Biqi, Nanuet, NY, UNITED STATES PAWYETH, Madison, NJ (U.S. corporation) PΙ US 2004110762 A1 20040610 US 2003-618044 A1 20030710 (10) ΑI Division of Ser. No. US 2000-751274, filed on 29 Dec 2000, GRANTED, Pat. RLI No. US 6638929 US 1999-240905P 19991229 (60) PRAI Utility DTFS APPLICATION Anne M. Rosenblum, Esq., Suite 212, 163 Delaware Avenue, Delmar, NY, LREP 12054 CLMN Number of Claims: 35 ECL Exemplary Claim: 1 DRWN No Drawings LN.CNT 8418 CAS INDEXING IS AVAILABLE FOR THIS PATENT. AB This invention provides compounds of formula 1, having the structure which are useful as inhibitors of protein tyrosine kinase and are antiproliferative agents. CAS INDEXING IS AVAILABLE FOR THIS PATENT. 348617-61-8P (preparation of substituted aromatic tricyclic compds. containing nicotinonitrile rings as protein kinase inhibitors) 348617-44-7P 348617-46-9P 348617-47-0P

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348617-50-5P 348617-51-6P 348617-52-7P
      348617-54-9P 348617-55-0P 348617-56-1P
      348617-58-3P 348617-62-9P 348617-71-0P
      348617-72-1P
         (preparation of substituted aromatic tricyclic compds. containing
nicotinonitrile
        rings as protein kinase inhibitors)
     348619-39-6P 348619-42-1P
         (preparation of substituted aromatic tricyclic compds. containing
nicotinonitrile
        rings as protein kinase inhibitors)
    ANSWER 2 OF 3 USPATFULL on STN 2001:229667 USPATFULL
L27
AN
TI
       Tricyclic protein kinase inhibitors
       Berger, Dan M., New City, NY, United States
IN
       Dutia, Minu D., West Nyack, NY, United States
DeMorin, Frenel F., Nanuet, NY, United States
       Boschelli, Diane H., New City, NY, United States
       Powell, Dennis W., Westchester, NY, United States
       Tsou, Hwei-Ru, New City, NY, United States Wissner, Allan, Ardsley, NY, United States Zhang, Nan, Eastchester, NY, United States
       Ye, Fei, Nanuet, NY, United States
       Wu, Biqi, Nanuet, NY, United States
       American Home Products Corporation, Madison, NJ, United States,
PA
       07940-0874 (U.S. corporation)
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PΙ
       US 2001051620
                            A1
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PRAI
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LREP
       MADISON, NJ, 07940-0874
       Number of Claims: 35
CLMN
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DRWN
LN.CNT 8432
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       This invention provides compounds of formula 1, having the structure
AB
       ##STR1##
       which are useful as inhibitors of protein tyro sine kinase and are
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CAS INDEXING IS AVAILABLE FOR THIS PATENT.
     348617-61-8P
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nicotinonitrile
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         (preparation of substituted aromatic tricyclic compds. containing
nicotinonitrile
        rings as protein kinase inhibitors)
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TT

348619-39-6P 348619-42-1P

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(preparation of substituted aromatic tricyclic compds. containing
nicotinonitrile
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L27 ANSWER 3 OF 3 USPAT2 on STN
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AN
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IN
       Berger, Dan M., New City, NY, United States
       Dutia, Minu D., West Nyack, NY, United States
       DeMorin, Frenel F., Nanuet, NY, United States
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       Wu, Biqi, Nanuet, NY, United States
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EXNAM Primary Examiner: Solola, Taopiq; Assistant Examiner: Sackey, Ebenezer
LREP
       Hogan, Jr., John W.
       Number of Claims: 16
CLMN
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       Exemplary Claim: 1
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CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       This invention provides compounds of formula 1, having the structure
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COPYRIGHT (c) 2004 Beilstein-Institut zur Foerderung der Chemischen Wissenschaften
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FILE RELOADED ON OCTOBER 20, 2002 FILE LAST UPDATED ON JUNE 15, 2004

FILE COVERS 1771 TO 2003. *** FILE CONTAINS 8,997,153 SUBSTANCES ***

>>> PLEASE NOTE: Reaction data and substance data are stored in separate documents and can not be searched together in one query.

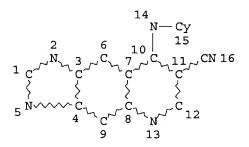
Reaction data for BEILSTEIN compounds may be displayed immediately with the display codes PRE (preparations) and REA (reactions). A substance answer set retrieved after the search for a chemical name, a molecular formula or a structure search for example can be restricted to compounds with available reaction information by concatenation with PRE/FA, REA/FA or more general with RX/FA. The BEILSTEIN Registry Number (BRN) is the link between a BEILSTEIN compound and belonging reactions. For more detailed reaction searches BRNs can be selected from substance answer sets and searched in the next step as reaction partner BRNs - Reactant (RX.RBRN) or Product BRN (RX.PBRN). After a search for reaction details substance documents associated with reactants or products may be retrieved by searching RX.PBRNs or RX.RBRNs as BRNs. <<<

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- * ARE BASED ON THE HIGHEST PRICE CATEGORY. THEREFORE; THESE
- * ESTIMATES MAY NOT REFLECT THE ACTUAL COSTS.
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wo dwgue stat 117

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STEREO ATTRIBUTES: NONE

L17 13 SEA FILE=BEILSTEIN SSS FUL L7 100.0% PROCESSED 22 ITERATIONS SEARCH TIME: 00.00.03

13 ANSWERS

=> d che ide ex 117 tob

L17 ANSWER 1 OF 13 BEILSTEIN COPYRIGHT 2004 BEILSTEIN MDL on STN

Beilstein Records (BRN): 9306940 Chemical Name (CN): 2-(2-morpholin-4-yl-ethylamino)-8-(3,4,5trimethoxy-phenylamino) -3H-imidazo<4,5g>quinoline-7-carbonitrile 2-(2-morpholin-4-yl-ethylamino)-8-(3,4,5-Autonom Name (AUN): trimethoxy-phenylamino) -3H-imidazo<4,5g>quinoline-7-carbonitrile Molec. Formula (MF): C26 H29 N7 O4 Molecular Weight (MW): 503.56 Lawson Number (LN): 30824, 30356, 15326, 3018, 289 Compound Type (CTYPE): heterocyclic Constitution ID (CONSID): 7859336 Tautomer ID (TAUTID): 8735609 Entry Date (DED): 2003/04/17 Update Date (DUPD): 2003/04/17

Field Availability:

Code	Name	Occurrence
======		
BRN	Beilstein Records	1
CN	Chemical Name	1
AUN	Autonomname	1
MF	Molecular Formula	1
FW	Formular Weight	1
LN	Lawson Number	5
FS	File Segment	1

CTYPE

```
Compound Type
                                                         1
     CONSID
               Constitution ID
                                                         1
     TAUTID
               Tautomer ID
                                                         1
              Entry Date
                                                         1
     UPD
              Update Date
                                                         1
     PHARM
               Pharmacological Data
                                                         5
   This substance also occurs in Reaction Documents:
     Code
              Name
                                               Occurrence
     Reaction Documents
     RXPRO
              Substance is Reaction Product
Reaction:
RX
     Reaction ID (.ID):
                                    9182059
     Reactant BRN (.RBRN):
                                    9294295, 1210530
     Reactant (.RCT):
                                    6,7-diamino-4-(3,4,5-trimethoxy-
                                    phenylamino) -quinoline-3-carbonitrile,
                                    4-(2-isothiocyanato-ethyl)-morpholine
     Product BRN (.PBRN):
                                    9306940
     Product (.PRO):
                                    2-(2-morpholin-4-yl-ethylamino)-8-(3,4,5-
                                    trimethoxy-phenylamino)-3H-imidazo<4,5-
                                    g>quinoline-7-carbonitrile
    No. of React. Details (.NVAR):
Reaction Details:
РX
    Reaction RID (.RID):
                                    9182059.1
    Reaction Classification (.CL): Multistage
    Nr. of Stages (.SNR):
    Stage 1
    Stage 2
    Reagent (.RGT):
                                    HqO
    Reference(s):
    1. Berger, Dan; Dutia, Minu; Powell, Dennis; Wu, Biqi; Wissner, Allan;
       DeMorin, Frenel; Weber, Jennifer; Boschelli, Frank,
       Bioorg.Med.Chem.Lett., CODEN: BMCLE8, 12(19), <2002>, 2761 - 2766;
       BABS-6374520
L17 ANSWER 2 OF 13 BEILSTEIN COPYRIGHT 2004 BEILSTEIN MDL on STN
    Beilstein Records (BRN):
                                    9304845
    Chemical Name (CN):
                                    8-(5-methoxy-2-methyl-phenylamino)-2-(2-
                                   morpholin-4-yl-ethylamino)-3H-imidazo<4,5-
                                   g>quinoline-7-carbonitrile
    Autonom Name (AUN):
                                    8-(5-methoxy-2-methyl-phenylamino)-2-(2-
                                   morpholin-4-yl-ethylamino)-3H-imidazo<4,5-
                                   g>quinoline-7-carbonitrile
    Molec. Formula (MF):
                                   C25 H27 N7 O2
    Molecular Weight (MW):
                                   457.53
    Lawson Number (LN):
                                   30824, 30356, 14902, 3018, 289
    Compound Type (CTYPE):
                                   heterocyclic
    Constitution ID (CONSID):
                                   7857619
    Tautomer ID (TAUTID):
                                   8734922
    Entry Date (DED):
                                   2003/04/17
    Update Date (DUPD):
                                   2003/04/17
```

Field Availability:

Code	Name	Occurrence
======	=======================================	=======================================
BRN	Beilstein Records	1
CN	Chemical Name	1
AUN	Autonomname	1
MF	Molecular Formula	1
FW	Formular Weight	1
LN	Lawson Number	5
CTYPE	Compound Type	1
CONSID	Constitution ID	1
TAUTID	Tautomer ID	1
ED	Entry Date	1
UPD	Update Date	1
PHARM	Pharmacological Data	5

This substance also occurs in Reaction Documents:

Code	Name	Occurrence
======	=======================================	=========
RX	Reaction Documents	1
RXPRO	Substance is Reaction Product	1

Reaction:

RX

```
Reaction ID (.ID):

Reactant BRN (.RBRN):

Reactant (.RCT):

6,7-diamino-4-(5-methoxy-2-methyl-phenylamino)-quinoline-3-carbonitrile,

4-(2-isothiocyanato-ethyl)-morpholine

Product BRN (.PBRN):

9304845

Product (.PRO):

8-(5-methoxy-2-methyl-phenylamino)-2-(2-morpholin-4-yl-ethylamino)-3H-imidazo<4,5-g>quinoline-7-carbonitrile

No. of React. Details (.NVAR):
```

```
Reaction Details:
ВX
```

Reaction RID (.RID): 9182058.1 Reaction Classification (.CL): Multistage

Nr. of Stages (.SNR):

Stage 1 Stage 2

Reagent (.RGT): Hg0

Reference(s):

 Berger, Dan; Dutia, Minu; Powell, Dennis; Wu, Biqi; Wissner, Allan; DeMorin, Frenel; Weber, Jennifer; Boschelli, Frank, Bioorg.Med.Chem.Lett., CODEN: BMCLE8, 12(19), <2002>, 2761 - 2766; BABS-6374520

L17 ANSWER 3 OF 13 BEILSTEIN COPYRIGHT 2004 BEILSTEIN MDL on STN

Beilstein Records (BRN): 9304516

Chemical Name (CN): 2-morpholin-4-ylmethyl-8-(3,4,5-trimethoxy-

phenylamino) -3H-imidazo<4,5-g>quinoline-7-

carbonitrile

Autonom Name (AUN): 2-morpholin-4-ylmethyl-8-(3,4,5-trimethoxy-

phenylamino) -3H-imidazo<4,5-q>quinoline-7-

carbonitrile C25 H26 N6 O4

Molec. Formula (MF): Molecular Weight (MW): 474.52

Lawson Number (LN): 30824, 30357, 15326, 289

Compound Type (CTYPE): heterocyclic Constitution ID (CONSID): 7857350 Tautomer ID (TAUTID): 8733004 Entry Date (DED): 2003/04/17 Update Date (DUPD): 2003/04/17

Field Availability:

```
Name
                                                      Occurrence
     BRN Bellstein ...
CN Chemical Name
               Beilstein Records
              Molecular Formula
     MF
               Formular Weight
     FW
     LN
               Lawson Number
     LN Lawson Number CTYPE Compound Type
     CONSID
               Constitution ID
     TAUTID Tautomer ID ED Entry Date
                                                               1
     UPD
                Update Date
                                                               1
     PHARM
                Pharmacological Data
   This substance also occurs in Reaction Documents:
     Code
              Name
                                                   Occurrence
     RX Reaction Documents
RXPRO Substance is Reaction Product
Reaction:
     Reaction ID (.ID):
                                        9177041
     Reactant BRN (.RBRN):
                                        9297462, 102549
     Reactant (.RCT):
                                       2-chloromethyl-8-(3,4,5-trimethoxy-
                                        phenylamino) -3H-imidazo<4,5-g>quinoline-7-
                                        carbonitrile, morpholine
     Product BRN (.PBRN):
                                        9304516
     Product (.PRO):
                                        2-morpholin-4-ylmethyl-8-(3,4,5-trimethoxy-
                                        phenylamino) -3H-imidazo<4,5-q>quinoline-7-
                                        carbonitrile
     No. of React. Details (.NVAR): 1
Reaction Details:
     Reaction RID (.RID):
                                        9177041.1
     Reaction Classification (.CL): Preparation
     Reference(s):
     1. Berger, Dan; Dutia, Minu; Powell, Dennis; Wu, Biqi; Wissner, Allan;
        DeMorin, Frenel; Weber, Jennifer; Boschelli, Frank,
        Bioorg.Med.Chem.Lett., CODEN: BMCLE8, 12(19), <2002>, 2761 - 2766;
        BABS-6374520
L17 ANSWER 4 OF 13 BEILSTEIN COPYRIGHT 2004 BEILSTEIN MDL on STN
     Beilstein Records (BRN): 9303785
                                        8-(4-chloro-5-methoxy-2-methyl-
     Chemical Name (CN):
                                        phenylamino) -3-(2-morpholin-4-yl-ethyl) -3H-
                                        imidazo<4,5-g>quinoline-7-carbonitrile
                                     8-(4-chloro-5-methoxy-2-methyl-
phenylamino)-3-(2-morpholin-4-yl-ethyl)-3H-
     Autonom Name (AUN):
    Molec. Formula (MF):

Molecular Weight (MW):

Lawson Number (LN):

Compound Type (CTYPE):

Constitution ID (CONSID):

Constitution (Z-morpholin-4-yl-ethyl)

imidazo<4,5-g>quinoline-7-carbonitrile

C25 H25 Cl N6 O2

476.96

30824, 30355, 14903, 3018, 289

heterocyclic

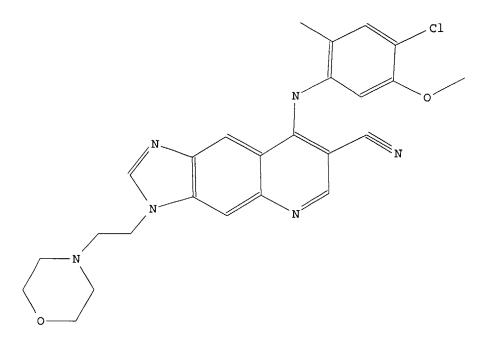
7856774
```

ВX

RX

Tautomer ID (TAUTID): Entry Date (DED): Update Date (DUPD):

8737736 2003/04/17 2003/04/17



Field Availability:

Code	Name	Occurrence
=======		=========
BRN	Beilstein Records	1
CN	Chemical Name	1
AUN	Autonomname	1
LSF	Linearized Structure Formula	1
MF	Molecular Formula	1
FW	Formular Weight	1
FBRN	Fragment BRN	2
LN	Lawson Number	5
FS	File Segment	1
CTYPE	Compound Type	1
CONSID	Constitution ID	1
TAUTID	Tautomer ID	1
ED	Entry Date	1
UPD	Update Date	1
PHARM	Pharmacological Data	5

This substance also occurs in Reaction Documents:

Code	Name	Occurrence
=======	=======================================	==========
RX	Reaction Documents	1
RXPRO	Substance is Reaction Product	1

Reaction:

```
Reaction ID (.ID):
                                     9192198
     Reactant BRN (.RBRN):
                                     9286129, 2831008
     Reactant (.RCT):
                                     8-chloro-3-(2-morpholin-4-yl-ethyl)-3H-
                                     imidazo<4,5-g>quinoline-7-carbonitrile,
                                     4-chloro-5-methoxy-2-methyl-aniline
     Product BRN (.PBRN):
                                     9303785
     Product (.PRO):
                                     8-(4-chloro-5-methoxy-2-methyl-
                                     phenylamino) -3-(2-morpholin-4-yl-ethyl) -3H-
                                     imidazo<4,5-g>quinoline-7-carbonitrile
     No. of React. Details (.NVAR):
Reaction Details:
ВX
     Reaction RID (.RID):
                                     9192198.1
     Reaction Classification (.CL): Preparation
     Reagent (.RGT):
                                     pyridine hydrochloride
     Solvent (.SOL):
                                     2-ethoxy-ethanol
     Other Conditions (.COND):
                                     Heating
     Reference(s):
     1. Berger, Dan; Dutia, Minu; Powell, Dennis; Wu, Biqi; Wissner, Allan;
        DeMorin, Frenel; Weber, Jennifer; Boschelli, Frank,
        Bioorg.Med.Chem.Lett., CODEN: BMCLE8, 12(19), <2002>, 2761 - 2766;
        BABS-6374520
L17 ANSWER 5 OF 13 BEILSTEIN COPYRIGHT 2004 BEILSTEIN MDL on STN
     Beilstein Records (BRN):
                                     9303647
     Chemical Name (CN):
                                     8-(2-bromo-4-chloro-5-methoxy-phenylamino) -
                                     3-(2-morpholin-4-yl-ethyl)-3H-imidazo<4,5-
                                     g>quinoline-7-carbonitrile
                                     8-(2-bromo-4-chloro-5-methoxy-phenylamino)-
     Autonom Name (AUN):
                                     3-(2-morpholin-4-yl-ethyl)-3H-imidazo<4,5-
                                     g>quinoline-7-carbonitrile
    Molec. Formula (MF):
                                     C24 H22 Br Cl N6 O2
    Molecular Weight (MW):
                                    541.83
    Lawson Number (LN):
                                    30824, 30355, 14894, 3018, 289
     Compound Type (CTYPE):
                                    heterocyclic
     Constitution ID (CONSID):
                                    7856662
     Tautomer ID (TAUTID):
                                    8737612
     Entry Date (DED):
                                    2003/04/17
    Update Date (DUPD):
                                    2003/04/17
```

Cl

Br

Field Availability:

Code	Name	Occurrence
======	=======================================	
BRN	Beilstein Records	1
CN	Chemical Name	1
AUN	Autonomname	1
MF	Molecular Formula	1
FW	Formular Weight	1
LN	Lawson Number	5
CTYPE	Compound Type	1
CONSID	Constitution ID	1
TAUTID	Tautomer ID	1
ED	Entry Date	1
UPD	Update Date	1
PHARM	Pharmacological Data	5

This substance also occurs in Reaction Documents:

Code	Name	Occurrence
======		=======================================
RX	Reaction Documents	1
RXPRO	Substance is Reaction Product	1

Reaction:

```
      Reaction ID (.ID):
      9214177

      Reactant BRN (.RBRN):
      9286129, 9044203

      Reactant (.RCT):
      8-chloro-3-(2-morpholin-4-yl-ethyl)-3H-imidazo<4,5-g>quinoline-7-carbonitrile,

      2-bromo-4-chloro-5-methoxy-phenylamine

      Product BRN (.PBRN):
      9303647

      Product (.PRO):
      8-(2-bromo-4-chloro-5-methoxy-phenylamino)-3-(2-morpholin-4-yl-ethyl)-3H-imidazo<4,5-</td>
```

q>quinoline-7-carbonitrile

No. of React. Details (.NVAR):

Reaction Details:

RX

Reaction RID (.RID): 9214177.1 Reaction Classification (.CL): Preparation

Reagent (.RGT): pyridine hydrochloride

Solvent (.SOL): 2-ethoxy-ethanol

Other Conditions (.COND): Heating

Reference(s):

 Berger, Dan; Dutia, Minu; Powell, Dennis; Wu, Biqi; Wissner, Allan; DeMorin, Frenel; Weber, Jennifer; Boschelli, Frank, Bioorg.Med.Chem.Lett., CODEN: BMCLE8, 12(19), <2002>, 2761 - 2766; BABS-6374520

L17 ANSWER 6 OF 13 BEILSTEIN COPYRIGHT 2004 BEILSTEIN MDL on STN

Beilstein Records (BRN): 9303524

Chemical Name (CN): 3-(2-morpholin-4-yl-ethyl)-8-(3,4,5-

trimethoxy-phenylamino)-3H-imidazo<4,5-

g>quinoline-7-carbonitrile

Autonom Name (AUN): 3-(2-morpholin-4-yl-ethyl)-8-(3,4,5-

trimethoxy-phenylamino)-3H-imidazo<4,5-

g>quinoline-7-carbonitrile

Molec. Formula (MF): C26 H28 N6 O4

Molecular Weight (MW): 488.55

Lawson Number (LN): 30824, 30355, 15326, 3018, 289

Compound Type (CTYPE): heterocyclic Constitution ID (CONSID): 7856543 Tautomer ID (TAUTID): 8737585

Entry Date (DED): 2003/04/17 Update Date (DUPD): 2003/04/17

Code	Name	Occurrence
======	=======================================	=======================================
BRN	Beilstein Records	1
CN	Chemical Name	1
AUN	Autonomname	1
MF	Molecular Formula	1
FW	Formular Weight	1
LN	Lawson Number	5
CTYPE	Compound Type	1
CONSID	Constitution ID	1
TAUTID	Tautomer ID	1
ED	Entry Date	1
UPD	Update Date	1
PHARM	Pharmacological Data	5

This substance also occurs in Reaction Documents:

Code	Name	Occurrence
	=======================================	========
RX	Reaction Documents	1
RXPRO	Substance is Reaction Product	1

Reaction:

RX

```
Reaction ID (.ID):

Reactant BRN (.RBRN):

Reactant (.RCT):

8-chloro-3-(2-morpholin-4-yl-ethyl)-3H-imidazo<4,5-g>quinoline-7-carbonitrile,
3,4,5-trimethoxy-aniline

Product BRN (.PBRN):

9303524

Product (.PRO):

3-(2-morpholin-4-yl-ethyl)-8-(3,4,5-trimethoxy-phenylamino)-3H-imidazo<4,5-g>quinoline-7-carbonitrile

No. of React. Details (.NVAR):
```

Reaction Details:

Reaction RID (.RID):

BABS-6374520

RX

```
Reaction Classification (.CL): Preparation
Reagent (.RGT): pyridine hydrochloride
Solvent (.SOL): 2-ethoxy-ethanol
Other Conditions (.COND): Heating
Reference(s):

1. Berger, Dan; Dutia, Minu; Powell, Dennis; Wu, Biqi; Wissner, Allan;
DeMorin, Frenel; Weber, Jennifer; Boschelli, Frank,
Bioorg.Med.Chem.Lett., CODEN: BMCLE8, 12(19), <2002>, 2761 - 2766;
```

9208412.1

L17 ANSWER 7 OF 13 BEILSTEIN COPYRIGHT 2004 BEILSTEIN MDL on STN

```
Beilstein Records (BRN): 9301278
Chemical Name (CN): 8-(2-chloro-5-methoxy-phenylamino)-3-(2-morpholin-4-yl-ethyl)-3H-imidazo<4,5-g>quinoline-7-carbonitrile
Autonom Name (AUN): 8-(2-chloro-5-methoxy-phenylamino)-3-(2-
```

```
morpholin-4-yl-ethyl)-3H-imidazo<4,5-
                                g>quinoline-7-carbonitrile
Molec. Formula (MF):
                                C24 H23 C1 N6 O2
Molecular Weight (MW):
                                462.94
Lawson Number (LN):
                                30824, 30355, 14893, 3018, 289
Compound Type (CTYPE):
                                heterocyclic
Constitution ID (CONSID):
                                7854680
Tautomer ID (TAUTID):
                                8736936
Entry Date (DED):
                                2003/04/17
Update Date (DUPD):
                                2003/04/17
```

Code	Name	Occurrence
=======		==============
BRN	Beilstein Records	1
CN	Chemical Name	1
AUN	Autonomname	1
MF	Molecular Formula	1
FW	Formular Weight	1
FBRN	Fragment BRN	2
LN	Lawson Number	5
FS	File Segment	1
CTYPE	Compound Type	1
CONSID	Constitution ID	1
TAUTID	Tautomer ID	1
ED	Entry Date	1
UPD	Update Date	1
PHARM	Pharmacological Data	5

This substance also occurs in Reaction Documents:

Code Name Occurrence

```
RX Reaction Documents
RXPRO Substance is Reaction Product
Reaction:
РX
     Reaction ID (.ID):
                                     9190208
     Reactant BRN (.RBRN):
                                     9286129, 2082193
     Reactant (.RCT):
                                     8-chloro-3-(2-morpholin-4-yl-ethyl)-3H-
                                     imidazo<4,5-g>guinoline-7-carbonitrile,
                                     2-chloro-5-methoxy-aniline
     Product BRN (.PBRN):
                                     9301278
     Product (.PRO):
                                     8-(2-chloro-5-methoxy-phenylamino)-3-(2-
                                     morpholin-4-yl-ethyl)-3H-imidazo<4,5-
                                     g>quinoline-7-carbonitrile
    No. of React. Details (.NVAR):
Reaction Details:
РX
     Reaction RID (.RID):
                                     9190208.1
    Reaction Classification (.CL): Preparation
    Reagent (.RGT):
                                     pyridine hydrochloride
    Solvent (.SOL):
                                     2-ethoxy-ethanol
    Other Conditions (.COND):
                                    Heating
    Reference(s):
     1. Berger, Dan; Dutia, Minu; Powell, Dennis; Wu, Biqi; Wissner, Allan;
        DeMorin, Frenel; Weber, Jennifer; Boschelli, Frank,
        Bioorg.Med.Chem.Lett., CODEN: BMCLE8, 12(19), <2002>, 2761 - 2766;
       BABS-6374520
L17 ANSWER 8 OF 13 BEILSTEIN COPYRIGHT 2004 BEILSTEIN MDL on STN
    Beilstein Records (BRN):
                                     9300518
    Chemical Name (CN):
                                     8-(5-methoxy-2-methyl-phenylamino)-3-(2-
                                     morpholin-4-yl-ethyl)-3H-imidazo<4,5-
                                     g>quinoline-7-carbonitrile
    Autonom Name (AUN):
                                     8-(5-methoxy-2-methyl-phenylamino)-3-(2-
                                     morpholin-4-yl-ethyl)-3H-imidazo<4,5-
                                     g>quinoline-7-carbonitrile
    Molec. Formula (MF):
                                    C25 H26 N6 O2
    Molecular Weight (MW):
                                    442.52
    Lawson Number (LN):
                                    30824, 30355, 14902, 3018, 289
    Compound Type (CTYPE):
                                   heterocyclic
    Constitution ID (CONSID):
                                    7854011
    Tautomer ID (TAUTID):
                                    8737502
                                   2003/04/17
    Entry Date (DED):
    Update Date (DUPD):
                                    2003/04/17
```

Code	Name	Occurrence
======	=======================================	=======
BRN	Beilstein Records	1
CN	Chemical Name	1
AUN	Autonomname	1
MF	Molecular Formula	1
FW	Formular Weight	1
LN	Lawson Number	5
CTYPE	Compound Type	1
CONSID	Constitution ID	1
TAUTID	Tautomer ID	1
ED	Entry Date	1
UPD	Update Date	1
PHARM	Pharmacological Data	5

This substance also occurs in Reaction Documents:

Code	Name	Occurrence
=======	=======================================	=========
RX	Reaction Documents	1
RXPRO	Substance is Reaction Product	1

Reaction:

```
Reaction ID (.ID): 9211265

Reactant BRN (.RBRN): 9286129, 774678

Reactant (.RCT): 8-chloro-3-(2-morpholin-4-yl-ethyl)-3H-
imidazo<4,5-g>quinoline-7-carbonitrile,
5-methoxy-2-methyl-aniline

Product BRN (.PBRN): 9300518

Product (.PRO): 8-(5-methoxy-2-methyl-phenylamino)-3-(2-
morpholin-4-yl-ethyl)-3H-imidazo<4,5-
```

g>quinoline-7-carbonitrile

No. of React. Details (.NVAR): 1

Reaction Details:

RX

Reaction RID (.RID): 9211265.1 Reaction Classification (.CL): Preparation

Reagent (.RGT): pyridine hydrochloride

Solvent (.SOL): 2-ethoxy-ethanol

Other Conditions (.COND): Heating

Reference(s):

 Berger, Dan; Dutia, Minu; Powell, Dennis; Wu, Biqi; Wissner, Allan; DeMorin, Frenel; Weber, Jennifer; Boschelli, Frank, Bioorg.Med.Chem.Lett., CODEN: BMCLE8, 12(19), <2002>, 2761 - 2766; BABS-6374520

L17 ANSWER 9 OF 13 BEILSTEIN COPYRIGHT 2004 BEILSTEIN MDL on STN

Beilstein Records (BRN): 9297462

Chemical Name (CN): 2-chloromethyl-8-(3,4,5-trimethoxy-

phenylamino) -3H-imidazo<4,5-g>quinoline-7-

carbonitrile

Autonom Name (AUN): 2-chloromethyl-8-(3,4,5-trimethoxy-

phenylamino) -3H-imidazo<4,5-q>quinoline-7-

carbonitrile

Molec. Formula (MF): C21 H18 C1 N5 O3

Molecular Weight (MW): 423.86

Lawson Number (LN): 30357, 15326, 289 Compound Type (CTYPE): heterocyclic

Compound Type (CTYPE): heterocyclic Constitution ID (CONSID): 7851478

Tautomer ID (TAUTID): 8728619
Entry Date (DED): 2003/04/17
Update Date (DUPD): 2003/04/17

Field Availability:

Code Name

Occurrence

```
BRN Beilstein Records
CN Chemical Name
     AUN
             Autonomname
           Molecular Formula
Formular Weight
Lawson Number
    MF
    FW
    LN
    FS
             File Segment
    CTYPE Compound Type
    CONSID
             Constitution ID
                                                       1
    TAUTID Tautomer ID ED Entry Date
                                                       1
                                                       1
    UPD
              Update Date
   This substance also occurs in Reaction Documents:
            Name
    RX Reaction Documents
             Substance is Reaction Reactant
    RXREA
    RXPRO Substance is Reaction Product
Reaction:
    Reaction ID (.ID):
                                  9205152
                            9294295, 605439
6,7-diamino-4-(3,4,5-trimethoxy-
phenylamino)-quinoline-3-carbonitrile,
    Reactant BRN (.RBRN):
    Reactant (.RCT):
                                  chloroacetyl chloride
    Product BRN (.PBRN):
                                   9297462
    Product (.PRO):
                                   2-chloromethyl-8-(3,4,5-trimethoxy-
                                  phenylamino) -3H-imidazo<4,5-q>quinoline-7-
                                   carbonitrile
    No. of React. Details (.NVAR): 1
Reaction Details:
    Reaction RID (.RID):
                                  9205152.1
    Reaction Classification (.CL): Multistage
    Nr. of Stages (.SNR):
    Stage 1
    Reagent (.RGT):
                                 Et2NPh
    Stage 2
    Reagent (.RGT):
                                  acetic acid
    Other Conditions (.COND): Heating
    Reference(s):
    1. Berger, Dan; Dutia, Minu; Powell, Dennis; Wu, Biqi; Wissner, Allan;
       DeMorin, Frenel; Weber, Jennifer; Boschelli, Frank,
       Bioorg.Med.Chem.Lett., CODEN: BMCLE8, 12(19), <2002>, 2761 - 2766;
       BABS-6374520
Reaction:
    Reaction ID (.ID):
                                  9177041
    Reactant BRN (.RBRN):
                               9297462, 102549
2-chloromethyl-8-(3,4,5-trimethoxy-
    Reactant (.RCT):
                                 phenylamino) -3H-imidazo<4,5-g>quinoline-7-
                                  carbonitrile, morpholine
    Product BRN (.PBRN):
                                  9304516
    Product (.PRO):
                                  2-morpholin-4-ylmethyl-8-(3,4,5-trimethoxy-
```

ВX

RX

phenylamino) -3H-imidazo<4,5-g>guinoline-7carbonitrile

No. of React. Details (.NVAR):

Reaction Details:

RX

Reaction RID (.RID): 9177041.1 Reaction Classification (.CL): Preparation

Reference(s):

1. Berger, Dan; Dutia, Minu; Powell, Dennis; Wu, Biqi; Wissner, Allan; DeMorin, Frenel; Weber, Jennifer; Boschelli, Frank, Bioorg.Med.Chem.Lett., CODEN: BMCLE8, 12(19), <2002>, 2761 - 2766; BABS-6374520

L17 ANSWER 10 OF 13 BEILSTEIN COPYRIGHT 2004 BEILSTEIN MDL on STN

Beilstein Records (BRN): 9292188 Chemical Name (CN): 8-(2-bromo-4-chloro-5-methoxy-phenylamino) -3H-imidazo<4,5-g>quinoline-7-carbonitrile Autonom Name (AUN): 8-(2-bromo-4-chloro-5-methoxy-phenylamino)-3H-imidazo<4,5-g>quinoline-7-carbonitrile

Molec. Formula (MF): C18 H11 Br Cl N5 O

Molecular Weight (MW): 428.67

Lawson Number (LN): 30355, 14894, 289 Compound Type (CTYPE): heterocyclic Constitution ID (CONSID): 7847049

Tautomer ID (TAUTID): 8725288 Entry Date (DED): 2003/04/17 Update Date (DUPD): 2003/04/17

Field Availability:

Code	Name	Occurrence
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BRN	Beilstein Records	1
CN	Chemical Name	1
AUN	Autonomname	1
MF	Molecular Formula	1
FW	Formular Weight	1

LN

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Lawson Number
     CTYPE
                Compound Type
     CONSID
                Constitution ID
                                                               1
     TAUTID
                Tautomer ID
     ED
                Entry Date
                                                               1
     UPD
                Update Date
                                                               1
     PHARM
                Pharmacological Data
   This substance also occurs in Reaction Documents:
               Name
                                                     Occurrence
     RX Reaction Documents
     RXPRO
               Substance is Reaction Product
Reaction:
RX
     Reaction ID (.ID):
                                       9214176
     Reaction ID (.ID).
Reactant BRN (.RBRN):
                                        9266116, 9044203
     Reactant (.RCT):
                                        8-chloro-3H-imidazo<4,5-g>quinoline-7-
                                        carbonitrile, 2-bromo-4-chloro-5-methoxy-
                                        phenylamine
     Product BRN (.PBRN):
                                        9292188
     Product (.PRO):
                                        8-(2-bromo-4-chloro-5-methoxy-phenylamino)-
                                        3H-imidazo<4,5-g>quinoline-7-carbonitrile
     No. of React. Details (.NVAR): 1
Reaction Details:
RX
     Reaction RID (.RID):
                                        9214176.1
     Reaction Classification (.CL): Preparation
                        pyridine hydrochloride
     Reagent (.RGT):
     Solvent (.SOL):
                                        2-ethoxy-ethanol
     Other Conditions (.COND): Heating
     Reference(s):
     1. Berger, Dan; Dutia, Minu; Powell, Dennis; Wu, Biqi; Wissner, Allan;
        DeMorin, Frenel; Weber, Jennifer; Boschelli, Frank,
        Bioorg.Med.Chem.Lett., CODEN: BMCLE8, 12(19), <2002>, 2761 - 2766;
        BABS-6374520
L17 ANSWER 11 OF 13 BEILSTEIN COPYRIGHT 2004 BEILSTEIN MDL on STN
     Beilstein Records (BRN):
                                        9291861
     Chemical Name (CN):
                                        8-(3,4,5-trimethoxy-phenylamino)-3H-
                                       imidazo<4,5-g>quinoline-7-carbonitrile
                                    imidazo<4,5-g>quinoline-7-carbonitri
8-(3,4,5-trimethoxy-phenylamino)-3H-
imidazo<4,5-g>quinoline-7-carbonitri
     Autonom Name (AUN):
                                      imidazo<4,5-g>quinoline-7-carbonitrile
     Molec. Formula (MF):
                                      C20 H17 N5 O3
    Molecular Weight (MW): 375.39

Lawson Number (LN): 30355, 15326,
Compound Type (CTYPE): heterocyclic
Constitution ID (CONSID): 7846818

Tautomer ID (TAUTID): 8725471

Entry Date (DED): 2003/04/17

Update Date (DUPD): 2003/04/17
                                   375.39
30355, 15326, 289
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Code	Name	Occurrence
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BRN	Beilstein Records	1
CN	Chemical Name	1
AUN	Autonomname	1
MF	Molecular Formula	1
FW	Formular Weight	1
LN	Lawson Number	3
CTYPE	Compound Type	1
CONSID	Constitution ID	1
TAUTID	Tautomer ID	1
ED	Entry Date	1
UPD	Update Date	1
NMR	Nuclear Magnetic Resonance	1
PHARM	Pharmacological Data	5

This substance also occurs in Reaction Documents:

Code	Name	Occurrence
=======	=======================================	========
RX	Reaction Documents	1
RXPRO	Substance is Reaction Product	1

Reaction:

RX

```
Reaction ID (.ID):

Reactant BRN (.RBRN):

Reactant (.RCT):

8-chloro-3H-imidazo<4,5-g>quinoline-7-carbonitrile, 3,4,5-trimethoxy-aniline

Product BRN (.PBRN):

9291861

Product (.PRO):

8-(3,4,5-trimethoxy-phenylamino)-3H-imidazo<4,5-g>quinoline-7-carbonitrile

No. of React. Details (.NVAR):
```

Reaction Details:

RX

Reaction RID (.RID): 9208408.1

Reaction Classification (.CL): Preparation

Reagent (.RGT): pyridine hydrochloride

Solvent (.SOL): 2-ethoxy-ethanol

Other Conditions (.COND): Heating

Reference(s):

 Berger, Dan; Dutia, Minu; Powell, Dennis; Wu, Biqi; Wissner, Allan; DeMorin, Frenel; Weber, Jennifer; Boschelli, Frank,

Bioorg.Med.Chem.Lett., CODEN: BMCLE8, 12(19), <2002>, 2761 - 2766;

BABS-6374520

Autonom Name (AUN):

L17 ANSWER 12 OF 13 BEILSTEIN COPYRIGHT 2004 BEILSTEIN MDL on STN

Beilstein Records (BRN): 9286966

Chemical Name (CN): 8-(2-chloro-5-methoxy-phenylamino)-3H-

imidazo<4,5-g>quinoline-7-carbonitrile
8-(2-chloro-5-methoxy-phenylamino)-3H-

imidazo<4,5-g>quinoline-7-carbonitrile
Molec. Formula (MF): C18 H12 Cl N5 O

Molec. Formula (MF): C18 H12 Molecular Weight (MW): 349.78

Lawson Number (LN): 30355, 14893, 289

Compound Type (CTYPE): heterocyclic

 Constitution ID (CONSID):
 7842596

 Tautomer ID (TAUTID):
 8723178

 Entry Date (DED):
 2003/04/17

 Update Date (DUPD):
 2003/04/17

Field Availability:

Code	Name	Occurrence
=======		=======================================
BRN	Beilstein Records	1
CN	Chemical Name	1
AUN	Autonomname	1
MF	Molecular Formula	1
FW	Formular Weight	1
LN	Lawson Number	3
CTYPE	Compound Type	1
CONSID	Constitution ID	1
TAUTID	Tautomer ID	1

1

ED

Entry Date

Update Date (DUPD):

```
UPD
               Update Date
                                                             1
     PHARM
                Pharmacological Data
   This substance also occurs in Reaction Documents:
     Code
              Name
     RX Reaction Documents
     RXPRO
               Substance is Reaction Product
Reaction:
RX
     Reaction ID (.ID):
                                      9190207
     Reactant BRN (.RBRN):
Reactant (.RCT):
                                      9266116, 2082193
                                      8-chloro-3H-imidazo<4,5-g>quinoline-7-
                                       carbonitrile, 2-chloro-5-methoxy-aniline
     Product BRN (.PBRN):
                                       9286966
     Product (.PRO):
                                       8-(2-chloro-5-methoxy-phenylamino)-3H-
                                       imidazo<4,5-g>quinoline-7-carbonitrile
     No. of React. Details (.NVAR):
Reaction Details:
RX
     Reaction RID (.RID):
                                       9190207.1
     Reaction Classification (.CL): Preparation
     Reagent (.RGT):
                                       pyridine hydrochloride
     Solvent (.SOL):
                                       2-ethoxy-ethanol
     Other Conditions (.COND):
                                      Heating
     Reference(s):
     1. Berger, Dan; Dutia, Minu; Powell, Dennis; Wu, Biqi; Wissner, Allan;
        DeMorin, Frenel; Weber, Jennifer; Boschelli, Frank,
        Bioorg.Med.Chem.Lett., CODEN: BMCLE8, 12(19), <2002>, 2761 - 2766;
        BABS-6374520
L17 ANSWER 13 OF 13 BEILSTEIN COPYRIGHT 2004 BEILSTEIN MDL on STN
     Beilstein Records (BRN): 9286576
Chemical Name (CN): 8-(2-bro
                                       8-(2-bromo-4-chloro-phenylamino)-3H-
                                       imidazo<4,5-g>quinoline-7-carbonitrile
     Autonom Name (AUN):
                                       8-(2-bromo-4-chloro-phenylamino)-3H-
                                      imidazo<4,5-g>quinoline-7-carbonitrile
                                 C17 H9 Br Cl N5
398.65
30355, 14133
    Molec. Formula (MF):

Molecular Weight (MW):

Lawson Number (LN):

Compound Type (CTYPE):

Constitution ID (CONSID):

Tautomer ID (TAUTID):

8724399

2003/04/17
     Molec. Formula (MF):
                                     2003/04/17
     Entry Date (DED):
```

2003/04/17

Code	Occurrence	
======	=======================================	===============
BRN	Beilstein Records	1
CN	Chemical Name	1
AUN	Autonomname	1
MF	Molecular Formula	1
FW	Formular Weight	1
LN	Lawson Number	2
CTYPE	Compound Type	1
CONSID	Constitution ID	1
TAUTID	Tautomer ID	1
ED	Entry Date	1
UPD	Update Date	1
PHARM	Pharmacological Data	5

This substance also occurs in Reaction Documents:

Code	Name	Occurrence
======	=======================================	
RX	Reaction Documents	1
RXPRO	Substance is Reaction Product	1

Reaction:

RX

```
Reaction ID (.ID):
                                9192082
Reactant BRN (.RBRN):
                                9266116, 2802563
Reactant (.RCT):
                                8-chloro-3H-imidazo<4,5-g>quinoline-7-
                                carbonitrile, 2-bromo-4-chloro-aniline
Product BRN (.PBRN):
                                9286576
Product (.PRO):
                                8-(2-bromo-4-chloro-phenylamino)-3H-
                                imidazo<4,5-g>quinoline-7-carbonitrile
No. of React. Details (.NVAR):
```

Reaction Details:

```
Reaction RID (.RID):
                                9192082.1
Reaction Classification (.CL):
                                Preparation
Reagent (.RGT):
                                pyridine hydrochloride
Solvent (.SOL):
                                2-ethoxy-ethanol
Other Conditions (.COND):
                                Heating
Reference(s):
```

 Berger, Dan; Dutia, Minu; Powell, Dennis; Wu, Biqi; Wissner, Allan; DeMorin, Frenel; Weber, Jennifer; Boschelli, Frank, Bioorg.Med.Chem.Lett., CODEN: BMCLE8, 12(19), <2002>, 2761 - 2766; BABS-6374520

=> b marpat

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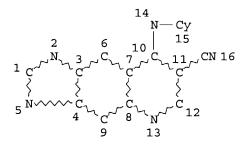
FILE CONTENT: 1988-PRESENT (VOL 141 ISS 04) (20040723/ED)

MOST RECENT CITATIONS FOR PATENTS FROM FIVE MAJOR ISSUING AGENCIES (COVERAGE TO THESE DATES IS NOT COMPLETE):

US 6747069 08 JUN 2004
DE 10351214 19 MAY 2004
EP 1424340 02 JUN 2004
JP 2004161736 10 JUN 2004
WO 2004052350 24 JUN 2004

Structure search limits have been raised. See HELP SLIMIT for the new, higher limits.

-> d que scat 120, L18 STR



NODE ATTRIBUTES:
DEFAULT MLEVEL IS ATOM
MLEVEL IS CLASS AT 15
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES: RING(S) ARE ISOLATED OR EMBEDDED NUMBER OF NODES IS 16

STEREO ATTRIBUTES: NONE

100.0% PROCESSED 7175 ITERATIONS 2 ANSWERS SEARCH TIME: 00.00.38

=> d bib abs 120 tot

L20 ANSWER 1 OF 2 MARPAT COPYRIGHT 2004 ACS on STN AN 136:37618 MARPAT

- *TI Preparation of substituted aromatic tricyclic compounds containing nicotinonitrile rings as protein kinase inhibitors
- Berger, Dan M.; Dutia, Minu D.; Demorin, Frenel F.; Boschelli, Diane H.; Powell, Dennis W.; Tsou, Hwei-ru; Wissner, Allan; Zhang, Nan; Ye, Fei; Wu, Biqi
- PA American Home Products Corporation, USA; Wyeth
- SO U.S. Pat. Appl. Publ., 107 pp. CODEN: USXXCO
- DT Patent
- LA English
- FAN. CNT 1

LAM.	CIVI I			
PATENT NO.		KIND	DATE	APPLICATION NO. DATE
ΡI	US 2001051620	A1	20011213	US 2000-751274 20001229
	US 6638929	B2	20031028	
	US 2004110762	A1	20040610	US 2003-618044 20030710
PRAI	US 1999-240905P	19991229		
	US 2000-751274	20001	.229	
GI				

- The title compds. I [Ar = (un) substituted cycloalkyl, pyridyl, pyrimidinyl, etc.; n = 0-1; X = NH, O, S, NR; R = alkyl; Y, Z = both carbon or N; A = (un) substituted benzo, pyrido, pyrimido, etc.] which are useful as inhibitors of protein tyrosine kinase and are antiproliferative agents, were prepared E.g., a 3-step synthesis of II which showed IC50 of 0.005 .mu.M against EGF-R kinase (recombinant enzyme), was given.
- L20 ANSWER 2 OF 2 MARPAT COPYRIGHT 2004 ACS on STN
- AN 135:92639 MARPAT
- TI Preparation of substituted aromatic tricyclic compounds containing nicotinonitrile rings as protein kinase inhibitors
- Berger, Dan M.; Dutia, Minu D.; Demorin, Frenel F.; Boschelli, Diane H.; Powell, Dennis W.; Tsou, Hwei-ru; Wissner, Allan; Zhang, Nan; Ye, Fei; Wu, Biqi
- PA American Home Products Corp., USA
- SO PCT Int. Appl., 377 pp.

CODEN: PIXXD2

- DT Patent
- LA English
- FAN.CNT 1

PATENT NO.			KI	ND :	DATE			APPLICATION NO.				o. 1	DATE				
									-								
PI WO 2001047892			Α	1	20010705			WO 2000-US35616					20001229				
	W:	ΑE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BY,	ΒZ,	CA,	CH,	CN,
		CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EE,	ES,	FI,	GB,	GD,	GE,	GH,	GM,	HR,
		HU,	ID,	IL,	IN,	IS,	JP,	KΕ,	KG,	ΚP,	KR,	ΚZ,	LC,	LK,	LR,	LS,	LT,
		LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NO,	NZ,	PL,	PT,	RO,	RU,
		WO 2001	WO 20010478 W: AE, CR, HU,	WO 2001047892 W: AE, AG, CR, CU, HU, ID,	WO 2001047892 A W: AE, AG, AL, CR, CU, CZ, HU, ID, IL,	WO 2001047892 A1 W: AE, AG, AL, AM, CR, CU, CZ, DE, HU, ID, IL, IN,	WO 2001047892 A1 2001 W: AE, AG, AL, AM, AT, CR, CU, CZ, DE, DK, HU, ID, IL, IN, IS,	WO 2001047892 A1 20010705 W: AE, AG, AL, AM, AT, AU, CR, CU, CZ, DE, DK, DM, HU, ID, IL, IN, IS, JP,	WO 2001047892 A1 20010705 W: AE, AG, AL, AM, AT, AU, AZ, CR, CU, CZ, DE, DK, DM, DZ, HU, ID, IL, IN, IS, JP, KE,	WO 2001047892 A1 20010705 W W: AE, AG, AL, AM, AT, AU, AZ, BA, CR, CU, CZ, DE, DK, DM, DZ, EE, HU, ID, IL, IN, IS, JP, KE, KG,	WO 2001047892 A1 20010705 WO 20 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, HU, ID, IL, IN, IS, JP, KE, KG, KP,	WO 2001047892 A1 20010705 WO 2000-US W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR,	WO 2001047892 A1 20010705 WO 2000-US356 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ,	WO 2001047892 A1 20010705 WO 2000-US35616 CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,	WO 2001047892 A1 20010705 WO 2000-US35616 2000 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK,	WO 2001047892 A1 20010705 WO 2000-US35616 20001229 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,	

SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM

RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG 20020925 EP 2000-988437 20001229 EP 1242382 Α1 AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR BR 2000-16878 20001229 BR 2000016878 Α 20021008 JP 2003519127 T2 20030617 JP 2001-549364 20001229 PRAI US 1999-473600 19991229 WO 2000-US35616 20001229 GΙ

$$\begin{bmatrix} \operatorname{CH}_2 \\ \operatorname{T} \\ \operatorname{CN} \end{bmatrix} = \begin{bmatrix} \operatorname{CN} \\ \operatorname{CN} \\ \operatorname{CN} \end{bmatrix}$$

AB The title compds. I [Ar = (un) substituted cycloalkyl, pyridyl, pyrimidinyl, etc.; n = 0-1; X = NH, O, S, NR; R = alkyl; Y, Z = both carbon or N; A = (un) substituted benzo, pyrido, pyrimido, etc.] which are useful as inhibitors of protein tyrosine kinase and are antiproliferative agents, were prepared E.g., a 3-step synthesis of II which showed IC50 of 0.005 .mu.M against EGF-R kinase (recombinant enzyme), was given.

RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

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